Claims:-

# 1. Use of a compound of formula I

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its salts, and pharmaceutically acceptable derivatives thereof, in the treatment of infections involving viruses of the *Pneumovirinae* sub-family, wherein

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A together with the atoms to which it is attached, forms an optionally substituted aromatic ring;

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linker B-C together with the atoms to which they are attached, forms an optionally substituted heterocyclic ring having from 5 to 8 ring atoms;

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 $R_1$  is selected from  $C_{1-12}$  alkyl,  $C_{2-12}$  alkenyl,  $C_{2-12}$  alkynyl,  $-(CH_2)_nC_{3-7}$  cycloalkyl,  $-(CH_2)_nC_{4-7}$  cycloalkenyl,  $-(CH_2)_n$  aryl,  $-(CH_2)_n$  aryl $C_{1-12}$  alkyl,  $-(CH_2)_n$  aryl $C_{2-12}$  alkynyl, and  $-(CH_2)_n$  heterocyclyl; n is 0-6 and the alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl and heterocyclyl groups are optionally substituted;

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 $R_2$  is selected from  $-CH_2R_3$ ,  $-C(Y)R_3$ ,  $-C(Y)OR_3$ ,  $-C(Y)N(R_4)R_3$ ,  $-C(Y)CH_2N(R_4)R_3$ ,  $-C(Y)CH_2SR_3$  and  $-S(O)_wR_5$ , where  $R_3$  is selected from hydrogen,  $C_{1-12}$  alkyl,  $C_{2-12}$  alkenyl,  $C_{2-12}$  alkynyl,  $-(CH_2)_mC_{3-7}$  cycloalkyl,  $-(CH_2)_mC_{4-7}$  cycloalkenyl,  $-(CH_2)_m$  aryl,  $-(CH_2)_m$  aryl $C_{1-12}$  alkyl,  $-(CH_2)_m$  aryl $C_{2-12}$  alkenyl,  $-(CH_2)_m$  aryl $C_{2-12}$  alkynyl and  $-(CH_2)_m$  heterocyclyl; and when  $R_2$  is  $-CH_2R_3$ , or  $-C(Y)R_3$ ,  $R_3$  may also be selected from  $-S-R_5$  and  $-O-R_5$ ; m is 0-6;  $R_4$  is hydrogen or  $C_{1-6}$  alkyl;  $R_5$  is  $C_{1-6}$  alkyl,  $C_{2-6}$  alkenyl,  $C_{2-6}$  alkynyl,  $C_{3-7}$  cycloalkyl,  $C_{4-7}$  cycloalkenyl, benzyl, aryl or heterocyclyl; w is 0, 1 or 2, and the alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl and heterocyclyl groups are optionally substituted;

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X and Y are independently selected from O, S and NR<sub>6</sub>, where R<sub>6</sub> is independently selected from hydrogen, lower alkyl, hydroxy and lower alkoxy.

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- 2. Use as defined in claim 1 wherein  $R_2$  is not an unsubstituted  $-C_{1-6}$ alkyl or unsubstituted  $-C(O)-C_{1-6}$ alkyl.
- 5 3. Use as defined in claim 1 wherein ring A is an optionally substituted aryl ring.
  - 4. Use as defined in claim 1 wherein ring A is an optionally substituted phenyl ring.
- 5. Use as defined in claim 1 wherein ring A is an optionally substituted heteroaryl ring.
  - 6. Use as defined in claim 1 wherein ring A together with the atoms to which it is attached, represents an optionally substituted pyridyl, pyridazinyl, pyrimidinyl, pyrazinyl, pyrrolyl, furyl, thienyl, imidazolyl, oxazolyl or isoxazolyl ring.
  - 7. Use as defined in claim 1 wherein ring A is an optionally substituted pyridyl, pyridazinyl, pyrimidinyl or pyrazinyl ring.
  - 8. Use as defined in claim 1 wherein ring A is optionally substituted pyridyl ring.
  - 9. Use as defined in claim 1 wherein ring A is optionally substituted with one or more substituents independently selected from halo, -NH<sub>2</sub>, NO<sub>2</sub>, C<sub>1-6</sub> alkyl, aryl and heterocyclyl, the aryl and heterocyclyl groups optionally substituted with halo, C<sub>1-6</sub> alkyl or halo substituted C<sub>1-6</sub> alkyl and, when ring A contains one or more ring nitrogens, the optional substituents include N-oxides of one or more of the ring nitrogens and pyridinium salts thereof.
- 10. Use as defined in claim 1 wherein ring A is optionally substituted with a substituent selected from halo, alkyl, C<sub>6</sub>H<sub>5</sub>- CH<sub>3</sub>-C<sub>6</sub>H<sub>4</sub>-, CF<sub>3</sub>-C<sub>6</sub>H<sub>4</sub>-, pyridyl, NO<sub>2</sub> and when ring A contains one or more ring nitrogens, the optional substituent also include an N-oxide form of a ring nitrogen, and pyridinium salts thereof.
  - 11. Use as defined in claim 1 wherein ring A is not substituted.
- 35 12. Use as defined in claim 1 of a compound of the formula IV

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its salts, N-oxides and pharmaceutically acceptable derivatives thereof, wherein B-C, X, R<sub>1</sub> and R<sub>2</sub> are as defined in claim 1.

- Use as defined in any one of claims 1 to 12, wherein R<sub>2</sub> is selected from -CH<sub>2</sub>R<sub>3</sub>, 13.  $-C(Y)R_3$ ,  $-C(Y)OR_3$ ,  $-C(Y)N(R_4)R_3$ ,  $-C(Y)CH_2N(R_4)R_3$ ,  $-C(Y)CH_2SR_3$  and  $-S(O)_wR_5$ , where R<sub>3</sub> is selected from hydrogen, -C<sub>1-12</sub>alkyl, -C<sub>2-12</sub>alkenyl, -C<sub>2-12</sub>alkynyl, -(CH<sub>2</sub>)<sub>m</sub>C<sub>3-</sub> 10 - $(CH_2)_mC_{4-7}$  cycloalkenyl, - $(CH_2)_m$ aryl, - $(CH_2)_m$ aryl $C_{1-12}$ 7cycloalkyl, -(CH<sub>2</sub>)<sub>m</sub>arylC<sub>2-12</sub>alkenyl, -(CH<sub>2</sub>)<sub>m</sub>arylC<sub>2-12</sub> alkynyl, -(CH<sub>2</sub>)<sub>m</sub>heterocyclyl, and when R<sub>2</sub> is -CH<sub>2</sub>R<sub>3</sub>, or -C(Y)R<sub>3</sub>, R<sub>3</sub> may also be selected from -S-R<sub>5</sub> and -O-R<sub>5</sub>; m is 0-6, R<sub>4</sub> is hydrogen or is C<sub>1-6</sub> alkyl, R<sub>5</sub> is selected from C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, C<sub>3-</sub> 7cycloalkyl, C<sub>4-7</sub> cycloalkenyl, benzyl, aryl and heterocyclyl; w is 0, 1 or 2, and the alkyl, 15 alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl and heterocyclyl groups are optionally substituted with one or more substituents selected from C<sub>1-6</sub> alkyl, C<sub>1-6</sub> alkoxy, C<sub>2-6</sub> alkenyl, C<sub>2-6</sub> alkynyl, halo, halo-C<sub>1-6</sub> alkyl (including CF<sub>3</sub>), hydroxy, mercapto, nitro, cyano, NH<sub>2</sub>, mono or di(C<sub>1-6</sub>alkyl) amino, phenyl, benzyl and heterocyclyl.
- Use as defined in claim 1 wherein  $R_2$  is  $-CH_2-R_3$ , and  $R_3$  is  $-(CH_2)_m$  aryl or  $-(CH_2)_m$  heterocyclyl and m is 0 to 3 and the aryl or heterocyclyl ring is optionally substituted.
  - 15. Use as defined in claim 1 wherein R<sub>2</sub> is -COR<sub>3</sub> and R<sub>3</sub> is aryl or heterocyclyl and is optionally substituted.
- 16. Use as defined in claim 14 or 15 wherein R<sub>3</sub> is optionally substituted phenyl, naphthyl, furyl, thienyl, pyrrolyl, H-pyrrolyl, pyrrolinyl, pyrrolidinyl, oxazolyl, oxadiazolyl, (including 1,2,3 and 1,2,4 oxadiazolyls) thiazolyl, isoxazolyl, furazanyl, isothiazolyl, pyrazolyl, pyrazolinyl, pyrazolidinyl, imidazolyl, imidazolinyl, triazolyl (including 1,2,3 and 1,3,4 triazolyls), tetrazolyl, thiadiazolyl (including 1,2,3 and 1,3,4 thiadiazolyls), pyridyl, pyrimidinyl, pyridazinyl, pyranyl, pyrazinyl, piperidinyl, 1,4-dioxanyl, morpholinyl, 1,4-dithianyl, thiomorpholinyl, piperazinyl, 1,3,5-trithianyl, triazinyl, 1H thieno[2,3-c]pyrazolyl, thieno[2,3-b]furyl, indolyl, isoindolyl, benzofuranyl, benzothienyl, benzoxazolyl, benzothiazolyl, benzisoxazolyl, benzisothiazolyl,

benzimidazolyl, indazolyl, isoquinolinyl, quinolinyl, quinoxalinyl, uridinyl, purinyl, cinnolinyl, phthalazinyl, quinazolinyl, quinoxalinyl, benzotriazinyl, naphthyridinyl or pteridinyl.

- Use as defined in claim 16, wherein R<sub>3</sub> is optionally substituted with one or more substituents selected from C<sub>1-6</sub> alkyl, C<sub>1-6</sub> alkoxy, C<sub>2-6</sub> alkenyl, C<sub>2-6</sub> alkynyl, halo, halo-C<sub>1-6</sub> alkyl (including CF<sub>3</sub>), hydroxy, mercapto, nitro, cyano, NH<sub>2</sub>, mono or di(C<sub>1-6</sub>alkyl) amino, phenyl, benzyl and heterocyclyl.
- 18. Use as defined in claim 1 wherein R<sub>2</sub> is -CON(H)R<sub>3</sub>, and R<sub>3</sub> is -(CH<sub>2</sub>)<sub>m</sub> aryl or (CH<sub>2</sub>)<sub>m</sub> heteroaryl and m is 0 to 2 and the aryl or heteroaryl ring is optionally substituted with one or more substituents independently selected from halo, lower alkyl, hydroxy, lower alkoxy and phenyl.
- 19. Use as defined in claim 1 wherein link -B-C- is an optionally substituted link of the formula -CH<sub>2</sub>-(CH<sub>2</sub>)<sub>z</sub>-, where z is 1-4.
  - 20. Use as defined in claim 19 wherein z is 1 or 2.
- 20 21. Use as defined in claim 1 wherein -B-C- is a linker of the formula -CH<sub>2</sub>CH<sub>2</sub>-.
  - 22. Use as defined in claim 1 wherein linker -B-C- is optionally substituted no more than three optional substituents, the substituents selected from halo, lower alkyl, hydroxy, lower alkoxy, phenyl and benzyl.
  - 23. Use as defined in claim 1 wherein linker -B-C- is not substituted.
    - 24. Use as defined in any one of claims 1 to 21 wherein X is oxygen or sulphur.
- 30 25. Use as defined in claim 1 wherein R<sub>1</sub> is an optionally substituted aryl or heterocyclyl group.
- 26. Use as defined in claim 1 wherein R<sub>1</sub> represents phenyl, thienyl, pyrrolyl, pyridyl ring or a -C<sub>1-6</sub> alkylphenyl group, the rings being optional substituted with halo, hydroxy, nitro, -NR'R" (where R' and R" are independently selected from hydrogen, lower alkyl and -C(O)R, where R is C<sub>1-6</sub> alkyl, phenyl or heterocyclyl), C<sub>1-12</sub>alkyl, phenyl and -O-R<sub>8</sub>, where R<sub>a</sub> is -C<sub>1-12</sub>alkyl, -C<sub>3-7</sub>cycloalkyl, -C<sub>1-12</sub>alkylC<sub>3-7</sub>cycloalkyl, phenyl or -C<sub>1-12</sub>alkylphenyl; and the C<sub>1-12</sub>alkyl, phenyl or R<sub>a</sub> group may be optionally substituted with halo, -CN, -NR'R", -CO<sub>2</sub>R or -CONR'R", where R, R' and R" are independently selected from hydrogen or lower alkyl.

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- Use as defined in claim 1 wherein R<sub>1</sub> is phenyl optionally substituted with a substituent selected from halo, -C<sub>1-6</sub>alkyl, -C<sub>1-6</sub>alkylhalo, -C<sub>1-6</sub>alkylCN, -OC<sub>1-6</sub>alkylCN, -OC<sub>1-6</sub>alkylCN, -OC<sub>1-6</sub>alkylCN, -OC<sub>1-6</sub>alkylC<sub>3-7</sub>cycloalkyl, -OC<sub>1-6</sub>alkylC<sub>6</sub>H<sub>5</sub>, -OC<sub>1-6</sub>alkylCO<sub>2</sub>NH<sub>2</sub>, -OC<sub>6</sub>H<sub>5</sub>, -OC<sub>6</sub>H<sub>4</sub>halo, -CF<sub>3</sub>, -OCF<sub>3</sub>, -NR'R" (where R' and R" are independently selected from hydrogen, -C(O)C<sub>1-6</sub>alkyl, -C(O)C<sub>6</sub>H<sub>5</sub>, -C(O)CH=CHCO<sub>2</sub>H, -C(O)C<sub>1-6</sub>alkylCO<sub>2</sub>H, -C(O)C<sub>1-6</sub>alkylCO<sub>2</sub>H, -C(O)C<sub>1-6</sub>alkylC<sub>6</sub>H<sub>4</sub>CH<sub>3</sub>, -C(O)C<sub>1-6</sub>alkylC<sub>6</sub>H<sub>4</sub>OCH<sub>3</sub> and -C(O)C<sub>1-6</sub>alkylC<sub>6</sub>H<sub>4</sub>halo), -CO<sub>2</sub>H, -CO<sub>2</sub>C<sub>1-6</sub>alkyl, -NO<sub>2</sub>, -OH, -C<sub>6</sub>H<sub>5</sub>, -C<sub>6</sub>H<sub>4</sub>C<sub>1-6</sub>alkyl, -C<sub>6</sub>H<sub>4</sub>halo and -OC(O)C<sub>1-6</sub>alkyl.
- 28. Use as defined in claim 1 wherein R<sub>1</sub> is phenyl substituted with halo, -OC<sub>1-6</sub>alkyl, -OC<sub>1-6</sub>alkylhalo, -OC<sub>1-6</sub>alkylCO<sub>2</sub>NH<sub>2</sub>, -OC<sub>1-6</sub>alkylCN, -OC<sub>1-6</sub>alkylC<sub>3-7</sub>cycloalkyl, -OC<sub>1-6</sub>alkylC<sub>6</sub>H<sub>5</sub> or -OC<sub>1-6</sub>alkylOCH<sub>3</sub>.
- 15 29. Use as defined in claim 1 wherein R<sub>1</sub> is 4-chlorophenyl.
  - 30. A method for the treatment of infections involving viruses of the *Pneumovirinae* sub-family by the inhibition of the virus's fusion processes by the administration of a therapeutically effective amount of a compound of formula I as defined in any one of claims 1 to 29, the salt or pharmaceutically acceptable derivatives thereof to a patient in need to treatment.
  - 31. A pharmaceutical formulation for the treatment of infections involving viruses of the *Pneumovirinae* sub-family comprising a compound of formula I as defined in any one of claims 1 to 29, the salt or pharmaceutically acceptable derivatives thereof.
    - 32. Use of a compound of formula I as defined in any one of claims 1 to 29, the salt or pharmaceutically acceptable derivatives thereof in the manufacture of a medicament for the treatment of infections involving viruses of the *Pneumovirinae* sub-family.
    - 33. A method for treating mammals infected with viruses of the *Pneumovirinae* subfamily, which comprises administering to the mammal a therapeutically effective amount of one or more of the compounds of formula I as defined in any one of claims 1 to 29, or pharmaceutically acceptable derivatives thereof.
  - 34. A method for preventing the infection of mammals with viruses of the *Pneumovirinae* sub-family, which comprises administering to the mammal a therapeutically effective amount of one or more of the compounds of formula I as defined in any one of claims 1 to 29, or pharmaceutically acceptable derivatives thereof.

- 35. The use or method according to any one of claims 1 to 34 in the treatment of infections involving viruses of the Pneumovirus and Metapneumovirus genus.
- The use or method according to any one of claims 1 to 34 in the treatment of 36. 5 respiratory syncytial virus (RSV).
  - The use or method according to any one of claims 1 to 34 in the treatment of 37. human RSV or human metapneumovirus.

#### 10 38. A compound of formula I

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Formula I

15 its salts, and pharmaceutically acceptable derivatives thereof, wherein

A together with the atoms to which it is attached, represents an optionally substituted phenyl, pyridyl, pyridazinyl, pyrimidinyl or pyrazinyl ring;

B-C is an optionally substituted link of the formula  $-CH_2-(CH_2)_z$ , where z is 1-4; 20

R<sub>1</sub> is selected from C<sub>1-12</sub> alkyl, C<sub>2-12</sub> alkenyl, C<sub>2-12</sub> alkynyl, -(CH<sub>2</sub>)<sub>n</sub>C<sub>3-7</sub> cycloalkyl, - $(CH_2)_nC_{4-7}$  cycloalkenyl, - $(CH_2)_n$  aryl, - $(CH_2)_n$  aryl $C_{1-12}$  alkyl, - $(CH_2)_n$  aryl $C_{2-12}$  alkenyl, - $(CH_2)_n$  ary  $iC_{2-12}$  alkynyl, and  $-(CH_2)_n$  heterocyclyl; n is 0-6 and the alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl and heterocyclyl groups are optionally substituted;

R<sub>2</sub> is selected from -CH<sub>2</sub>R<sub>3</sub>, -C(Y)R<sub>3</sub>, -C(Y)OR<sub>3</sub>, -C(Y)N(R<sub>4</sub>)R<sub>3</sub> and -S(O)<sub>w</sub>R<sub>5</sub>, where R<sub>3</sub> is selected from hydrogen, C<sub>1-12</sub> alkyl, C<sub>2-12</sub> alkenyl, C<sub>2-12</sub> alkynyl, -(CH<sub>2</sub>)<sub>m</sub>C<sub>3-7</sub> cycloalkyl, - $(CH_2)_mC_{4-7}$  cycloalkenyl, - $(CH_2)_m$  aryl, - $(CH_2)_m$  aryl $C_{1-12}$  alkyl, - $(CH_2)_m$  aryl $C_{2-12}$  alkenyl, -(CH<sub>2</sub>)<sub>m</sub> arylC<sub>2-12</sub> alkynyl and -(CH<sub>2</sub>)<sub>m</sub> heterocyclyl; and when R<sub>2</sub> is -CH<sub>2</sub>R<sub>3</sub>, or -C(Y)R<sub>3</sub>. 30 R<sub>3</sub> may also be selected from -S-R<sub>5</sub> and -O-R<sub>5</sub>; m is 0-6; R<sub>4</sub> is hydrogen or C<sub>1-6</sub> alkyl; R<sub>5</sub> is C<sub>1-6</sub> alkyl, C<sub>2-6</sub> alkenyl, C<sub>2-6</sub> alkynyl, C<sub>3-7</sub> cycloalkyl, C<sub>4-7</sub> cycloalkenyl, benzyl, aryl or heterocyclyl; w is 0, 1 or 2, and the alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl and heterocyclyl groups are optionally substituted,

X and Y are independently selected from O, S and NR<sub>6</sub>, where R<sub>6</sub> is independently selected from hydrogen, lower alkyl, hydroxy and lower alkoxy;

with the provisos that when A is phenyl and R<sub>1</sub> is 4-chlorophenyl or unsubstituted phenyl

- R<sub>3</sub> is not unsubstituted cyclopropyl, halomethyl, unsubstituted phenyl or phenyl with only halo, -CH<sub>3</sub> and/or -OCH<sub>3</sub> substituents when R<sub>2</sub> is COR<sub>3</sub>;
  - (ii) R<sub>3</sub> is not unsubstituted phenyl or phenyl with only halo, -CH<sub>3</sub>, -OCH<sub>3</sub> and/or -C(O)OCH<sub>2</sub>CH<sub>3</sub> substituents when R<sub>2</sub> is C(O)NHR<sub>3</sub>;
- (iii) R<sub>3</sub> is not unsubstituted phenyl or phenyl with only halo, -CH<sub>3</sub>, -OCH<sub>3</sub> and/or -C(O)OCH<sub>2</sub>CH<sub>3</sub> substituents when R<sub>2</sub> is C(S)NHR<sub>3</sub>;

### and with the provisos

- (iv) when A is phenyl and R<sub>2</sub> is CH<sub>2</sub>R<sub>3</sub>, R<sub>3</sub> is not hydrogen, unsubstituted C<sub>1-6</sub> alkyl or C<sub>1-6</sub> alkyl only substituted with NH<sub>2</sub>, mono or di C<sub>1-6</sub> alkyl amino groups;
- 15 (v) when A is phenyl and R<sub>1</sub> is 4-methoxyphenyl, R<sub>2</sub> is not CHO;
  - when A is phenyl and R<sub>1</sub> is phenyl optionally substituted with only halo, C<sub>1-6</sub> alkyl and / or C<sub>1-6</sub> alkoxy and R<sub>2</sub> is COR<sub>3</sub>, R<sub>3</sub> is not methylene substituted with NH<sub>2</sub>, mono or di C<sub>1-6</sub> alkyl amino, N-piperidinyl or N-morpholinyl;
- (vii) when A is phenyl and R<sub>1</sub> is 3-CH<sub>3</sub>,4-CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>NHC(O)CH<sub>2</sub>O-phenyl, R<sub>2</sub> is not -S(O)<sub>2</sub>CH<sub>2</sub>SO<sub>2</sub>CH<sub>3</sub>, -CHO, -COCH<sub>2</sub>CH<sub>3</sub>, -CH<sub>2</sub>CH<sub>2</sub>OH, -CH<sub>2</sub>CH<sub>2</sub>OCH<sub>3</sub>, -CH<sub>2</sub>CO<sub>2</sub>C(CH<sub>3</sub>)<sub>3</sub> or C<sub>1-6</sub> alkyl;
  - (viii) when A is pyridyl and R<sub>1</sub> is 3-CH<sub>3</sub>,4-CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>NHC(O)CH<sub>2</sub>O-phenyl, R<sub>2</sub> is not CH<sub>3</sub>.
- 25 39. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, with the proviso that when ring A is phenyl
  - (i) R<sub>3</sub> is not hydrogen or optionally substituted C<sub>1-6</sub> alkyl when R<sub>2</sub> is -CH<sub>2</sub>R<sub>3</sub> or -COR<sub>3</sub>;
- R<sub>3</sub> is not (CH<sub>2</sub>)<sub>m</sub>heterocyclyl where m is 1 or 2 and the heterocyclyl ring is piperidinyl, morpholinyl, pyrrolidinyl, piperazinyl, thiomorpholinyl when R<sub>2</sub> is COR<sub>3</sub> and R<sub>1</sub> is 4-chlorophenyl, 4-methoxyphenyl or unsubstituted phenyl;
  - (iii) R<sub>2</sub> is not benzyl; and with the proviso

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- (iv) R<sub>2</sub> is not -CH<sub>3</sub> when A is pyridyl.
- 40. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, when A is phenyl and  $R_2$  is  $-CH_2R_3$  or  $-C(O)R_3$ , and  $R_3$  is selected from  $C_{7-12}$  alkyl,  $C_{2-12}$  alkenyl,  $C_{2-12}$  alkynyl,  $-(CH_2)_mC_{3-7}$  cycloalkyl,  $-(CH_2)_mC_{4-7}$  cycloalkenyl,  $-(CH_2)_m$  aryl,  $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  heterocyclyl,  $-(CH_2)_m$  and  $-(CH_2)_m$  aryl $-(CH_2)_m$  heterocyclyl,  $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  heterocyclyl,  $-(CH_2)_m$  aryl $-(CH_2)_m$  heterocyclyl,  $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  heterocyclyl,  $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  aryl $-(CH_2)_m$  heterocyclyl,  $-(CH_2)_m$  aryl $-(CH_2)_m$

- 41. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein ring A is optionally substituted with one or more substituents independently selected from halo, -NH<sub>2</sub>, NO<sub>2</sub>, C<sub>1-6</sub> alkyl, aryl and heterocyclyl, the aryl and heterocyclyl groups optionally substituted with halo, C<sub>1-6</sub>alkyl or halo substituted C<sub>1-6</sub> alkyl and, when ring A contains one or more ring nitrogens, the optional substituents include N-oxides of one or more of the ring nitrogens.
- 42. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein ring A is optionally substituted with a substituent selected from halo, alkyl, C<sub>6</sub>H<sub>5</sub>- CH<sub>3</sub>-C<sub>6</sub>H<sub>4</sub>-, CF<sub>3</sub>-C<sub>6</sub>H<sub>4</sub>-, pyridyl, NO<sub>2</sub> and when ring A contains one or more ring nitrogens, the optional substituent also include an N-oxide form of a ring nitrogen.
- 15 43. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein ring A is not substituted.
- The compound as defined in claim 38, the salt or pharmaceutically acceptable 44. derivative thereof, wherein R<sub>2</sub> is selected from -CH<sub>2</sub>R<sub>3</sub>, -C(Y)R<sub>3</sub>, -C(Y)OR<sub>3</sub>, -C(Y)N(R<sub>4</sub>)R<sub>3</sub>, -C(Y)CH<sub>2</sub>N(R<sub>4</sub>)R<sub>3</sub>, -C(Y)CH<sub>2</sub>SR<sub>3</sub> and -S(O)<sub>w</sub>R<sub>5</sub>, where R<sub>3</sub> is selected from 20 hydrogen,  $-C_{1-12}$ alkyl,  $-C_{2-12}$ alkenyl,  $-C_{2-12}$ alkynyl,  $-(CH_2)_mC_{3-7}$ cycloalkyl,  $-(CH_2)_mC_{4-7}$ cycloalkenyl, -(CH<sub>2</sub>)<sub>m</sub>aryl, -(CH<sub>2</sub>)<sub>m</sub>arylC<sub>1-12</sub> alkyl, -(CH<sub>2</sub>)<sub>m</sub>arylC<sub>2-12</sub>alkenyl, (CH<sub>2</sub>)<sub>m</sub>arylC<sub>2-12</sub> alkynyl, -(CH<sub>2</sub>)<sub>m</sub>heterocyclyl, and when R<sub>2</sub> is -CH<sub>2</sub>R<sub>3</sub>, or -C(Y)R<sub>3</sub>, R<sub>3</sub> may also be selected from -S-R<sub>5</sub> and -O-R<sub>5</sub>; m is 0-6, R<sub>4</sub> is hydrogen or is C<sub>1-6</sub> alkyl, R<sub>5</sub> is selected from C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, C<sub>3-7</sub>cycloalkyl, C<sub>4-7</sub> cycloalkenyl, benzyl, 25 aryl and heterocyclyl; w is 0, 1 or 2, and the alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl and heterocyclyl groups are optionally substituted with one or more substituents selected from C<sub>1-6</sub> alkyl, C<sub>1-6</sub> alkoxy, C<sub>2-6</sub> alkenyl, C<sub>2-6</sub> alkynyl, halo, halo-C<sub>1-6</sub> alkyl (including CF<sub>3</sub>), hydroxy, mercapto, nitro, cyano, NH<sub>2</sub>, mono or di(C<sub>1-6</sub>alkyl) amino, phenyl, benzyl and heterocyclyl, the substituents being optionally substituted. **30** 
  - 45. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein  $R_2$  is  $-CH_2-R_3$ , and  $R_3$  is  $-(CH_2)_m$  aryl or  $-(CH_2)_m$  heterocyclyl and m is 0 to 3 and the aryl or heterocyclyl ring is optionally substituted.
  - 46. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein R<sub>2</sub> is -COR<sub>3</sub> and R<sub>3</sub> is aryl or heterocyclyl and is optionally substituted.

- 47. The compound as defined in claim 45 or 46, the salt or pharmaceutically acceptable derivative thereof, wherein R<sub>3</sub> is optionally substituted phenyl, naphthyl, furyl, thienyl, pyrrolyl, H-pyrrolyl, pyrrolinyl, pyrrolidinyl, oxazolyl, oxadiazolyl, (including 1,2,3 and 1,2,4 oxadiazolyls) thiazolyl, isoxazolyl, furazanyl, isothiazolyl, pyrazolyl, pyrazolinyl, pyrazolidinyl, imidazolyl, imidazolinyl, triazolyl (including 1,2,3 and 1,3,4 triazolyls), tetrazolyl, thiadiazolyl (including 1,2,3 and 1,3,4 thiadiazolyls), pyridyl, pyrimidinyl, pyridazinyl, pyranyl, pyrazinyl, piperidinyl, 1,4-dioxanyl, morpholinyl, 1,4-dithianyl, thiomorpholinyl, piperazinyl, 1,3,5-trithianyl, triazinyl, 1H thieno[2,3-c]pyrazolyl, thieno[2,3-b]furyl, indolyl, isoindolyl, benzofuranyl, benzothienyl, benzoxazolyl, benzisoxazolyl, benzisothiazolyl, benzisothiazolyl, indazolyl, isoquinolinyl, quinoxalinyl, quinoxalinyl, uridinyl, purinyl, cinnolinyl, phthalazinyl, quinazolinyl, quinoxalinyl, naphthyridinyl or pteridinyl.
- 48. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein R<sub>3</sub> is optionally substituted with one or more substituents selected from C<sub>1-6</sub> alkyl, C<sub>1-6</sub> alkoxy, C<sub>2-6</sub> alkenyl, C<sub>2-6</sub> alkynyl, halo, halo-C<sub>1-6</sub> alkyl (including CF<sub>3</sub>), hydroxy, mercapto, nitro, cyano, NH<sub>2</sub>, mono or di(C<sub>1-6</sub>alkyl) amino, phenyl, benzyl and heterocyclyl, the phenyl, benzyl and heterocyclyl groups being optionally substituted.
  - 49. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein  $R_2$  is  $-CON(H)R_3$ , and  $R_3$  is  $-(CH_2)_m$  aryl or  $-(CH_2)_m$  heteroaryl and m is 0 to 2 and the aryl or heteroaryl ring is optionally substituted with one or more substituents independently selected from halo, lower alkyl, hydroxy, lower alkoxy and phenyl.
  - 50. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein z is 1 or 2.
- 30 51. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein -B-C- is a linker of the formula -CH<sub>2</sub>CH<sub>2</sub>-.
  - 52. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein the linker -B-C- is optionally substituted no more than three optional substituents, the substituents selected from halo, lower alkyl, hydroxy, lower alkoxy, phenyl and benzyl.
    - 53. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein the linker -B-C- is not substituted.

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- 54. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein X is oxygen or sulphur.
- 55. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein X is oxygen.
  - 56. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein R<sub>1</sub> is an optionally substituted aryl or heterocyclyl group.
- The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein R<sub>1</sub> represents phenyl, thienyl, pyrrolyl, pyridyl ring or a -C<sub>1-6</sub> alkylphenyl group, the rings being optional substituted with halo, hydroxy, nitro, -NR'R" (where R' and R" are independently selected from hydrogen, lower alkyl and -C(O)R, where R is C<sub>1-6</sub> alkyl, phenyl or heterocyclyl), C<sub>1-12</sub>alkyl, phenyl and -O-R<sub>a</sub>, where R<sub>a</sub> is -
- $C_{1-12}$ alkyl,  $-C_{3-7}$ cycloalkyl,  $-C_{1-12}$ alkyl $C_{3-7}$ cycloalkyl, phenyl or  $-C_{1-12}$ alkylphenyl; and the  $C_{1-12}$ alkyl, phenyl or  $R_a$  group may be optionally substituted with halo, -CN, -NR'R'',  $-CO_2R$  or -CONR'R'', where R, R' and R'' are independently selected from hydrogen or lower alkyl.
- 58. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein R<sub>1</sub> is phenyl optionally substituted with a substituent selected from halo, -C<sub>1-6</sub>alkyl, -C<sub>1-6</sub>alkylhalo, -C<sub>1-6</sub>alkylCN, -OC<sub>1-6</sub>alkyl, -OC<sub>1-6</sub>alkylhalo, -OC<sub>1-6</sub>alkylCO<sub>2</sub>NH<sub>2</sub>, -OC<sub>1-6</sub>alkylCN, -OC<sub>1-6</sub>alkylC<sub>3-7</sub>cycloalkyl, -OC<sub>1-6</sub>alkylC<sub>6</sub>H<sub>5</sub>, -OC<sub>1-6</sub>alkylOCH<sub>3</sub>, -OC<sub>6</sub>H<sub>5</sub>, -OC<sub>6</sub>H<sub>4</sub>halo, -CF<sub>3</sub>, -OCF<sub>3</sub>, -NR'R" (where R' and R" are
- independently selected from hydrogen, -C(O)C<sub>1-6</sub>alkyl, -C(O)C<sub>6</sub>H<sub>5</sub>, -C(O)CH=CHCO<sub>2</sub>H<sub>7</sub>, -C(O)C<sub>1-6</sub>alkylCO<sub>2</sub>H<sub>7</sub>, -C(O)C<sub>1-6</sub>alkylCO<sub>2</sub>CH<sub>3</sub>, -C(O)C<sub>1-6</sub>alkylC<sub>6</sub>H<sub>5</sub>, -C(O)C<sub>1-6</sub>alkylC<sub>6</sub>H<sub>4</sub>CH<sub>3</sub>, -C(O)C<sub>1-6</sub>alkylC<sub>6</sub>H<sub>4</sub>OCH<sub>3</sub> and -C(O)C<sub>1-6</sub>alkylC<sub>6</sub>H<sub>4</sub>halo), -CO<sub>2</sub>H<sub>1</sub>, -CO<sub>2</sub>C<sub>1-6</sub>alkyl, -NO<sub>2</sub>, -OH<sub>1</sub>, -C<sub>6</sub>H<sub>4</sub>C<sub>1-6</sub>alkyl, -C<sub>6</sub>H<sub>4</sub>halo and -OC(O)C<sub>1-6</sub>alkyl.
- 30 59. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein R<sub>1</sub> is halo-phenyl.
  - 60. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivative thereof, wherein  $R_1$  is 4-chlorophenyl.
  - 61. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivatives thereof, wherein A is an optionally substituted phenyl ring.

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- 62. The compound as defined in claim 38, the salt or pharmaceutically acceptable derivatives thereof, wherein  $R_2$  is  $C(O)-R_3$  and  $R_3$  is  $-(CH_2)_m$ -aryl or  $(CH_2)_m$ -heteroaryl, where m is 0 to 6, and the aryl or heteroaryl group is optionally substituted.
- 5 63. The compound as defined in claim 38 of the formula IV

Formula IV

- wherein R<sub>1</sub>, R<sub>2</sub>, X and -B-C- are as defined in claim 38, and the N-oxide form and pyridium salt thereof.
- 64. The compound as defined in claim 63, and the N-oxide form and pyridium salt thereof, wherein R<sub>2</sub> is C(O)R<sub>3</sub> and R<sub>3</sub> is -(CH<sub>2</sub>)<sub>m</sub>-aryl or (CH<sub>2</sub>)<sub>m</sub>-heteroaryl, where m is 0 to 6, and the aryl or heteroaryl group is optionally substituted.
  - 65. A compound disclosed in table 2 or 3.
- 66. A pharmaceutical formulation for the treatment of infections involving viruses of Pneumovirinae sub-family comprising a compound of formula I as defined in any one of claims 38 to 65, the salt or pharmaceutically acceptable derivative thereof.

PCT/AU2004/001830

# AP20 Rec'd PCT/PTO 23 JUN 2006 AMENDED CLAIMS

[received by the International Bureau on 10 May 2005 (10.05.05); new claims 67-82 added; remaining claims unchanged (3 pages)]

67. A compound of formula

and salts thereof, wherein

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the pyridyl ring is optionally substituted;

B-C is an optionally substituted linker of the formula  $-CH_2-(CH_2)_z$ , where z is 1-4;

- R<sub>1</sub> is selected from  $C_{1-12}$  alkyl,  $C_{2-12}$  alkenyl,  $C_{2-12}$  alkynyl,  $-(CH_2)_nC_{3-7}$  cycloalkyl,  $-(CH_2)_nC_{4-7}$  cycloalkenyl,  $-(CH_2)_n$  aryl,  $-(CH_2)_n$  aryl $C_{1-12}$  alkyl,  $-(CH_2)_n$  aryl $C_{2-12}$  alkenyl, aryl $C_{2-12}$  alkynyl, and  $-(CH_2)_n$  heterocyclyl; n is 0-6 and the alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl and heterocyclyl groups are optionally substituted;
- 15 X is selected from O, S and NR<sub>6</sub>, where R<sub>6</sub> is independently selected from hydrogen, lower alkyl, hydroxy and lower alkoxy;

with the proviso that when -B-C- is -CH<sub>2</sub>CH(CH(CH<sub>3</sub>)<sub>2</sub>)-, R<sub>1</sub> is not 3-CH<sub>3</sub>,4-CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>NHC(O)CH<sub>2</sub>O-phenyl-.

- 68. The compound as defined in claims 67 and salts thereof, wherein the pyridyl ring is optionally substituted with one or more substituents independently selected from halo,  $-NH_2$ ,  $-NO_2$ ,  $-C_{1-6}$ alkyl, aryl and heterocyclyl, the aryl and heterocyclyl groups optionally substituted with halo,  $C_{1-6}$ alkyl or halo substituted  $C_{1-6}$  alkyl, and the ring nitrogen of the pyridyl ring may optionally be an N-oxide.
- 69. The compound as defined in claim 67 and salts thereof, wherein the pyridyl ring is optionally substituted with a substituent selected from halo, alkyl,  $C_6H_5$ -,  $CH_3$ - $C_6H_4$ -,  $CF_3$ - $C_6H_4$ -, pyridyl and  $NO_2$ , and the ring nitrogen of the pyridyl ring may optionally be an N-oxide.
- 70. The compound as defined claim 67 and salts thereof, wherein the pyridyl ring is not substituted.
- 71. The compound as defined in claim 67 and salts thereof, wherein the linker -B-C- is as defined in any one of claims 21 to 23.

# **AMENDED SHEET (ARTICLE 19)**

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- 72. The compound as defined in claim 67 and salts thereof, wherein X is oxygen or sulphur.
- 5 73. The compound as defined in claim 67 and salts thereof, wherein X is oxygen.
  - 74. The compound as defined in claim 67 and salts thereof, wherein  $R_1$  is as defined in any one of claims 25 to 29.
- 10 75. A compound of formula

and salts thereof, wherein the pyridyl ring is optionally substituted and  $R_1$  and X are as defined in Claim 67, with the proviso that  $R_1$  is not 4-chlorophenyl.

76. A compound of the formula

and salts thereof, wherein the fused pyridazinyl ring is optionally substituted and R<sub>1</sub> and X are as defined in Claim 67, with the proviso that R<sub>1</sub> is not phenyl, 4-chlorophenyl or 4-methoxyphenyl.

77. A compound of any one of the formula

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and salts thereof, wherein the fused pyridyl, pyrazinyl, pyridazinyl or pyrimidinyl ring is optionally substituted and R<sub>1</sub> and X are as defined in Claim 67.

5 78. Use of a compound of formula III,

Formula III

and salts thereof, wherein R<sub>1</sub>, ring A, -B-C- and X are as defined in claim 38, as an intermediate for the production of a compound of formula I as defined in claim 38.

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- 79. A method of separating the enantiomers of a compound of formula III by forming diastereomeric salts of the compounds using an enantiomerically enriched chiral hydrogen phosphate.
- 15 80. A method of separating the enantiomers of a compound as defined in claim 67 by forming diastereomeric salts of the compound using an enantiomerically enriched chiral hydrogen phosphate.
  - 81. The compound as defined in claim 38 in a substantially pure optically active form.

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82. The compound as defined in claim 67, 75, 76 or 77 in a substantially pure optically active form.